Passion for Innovation. Compassion for Patients.™



ENHERTU® Business Briefing

DAIICHI SANKYO CO., LTD.

Sunao Manabe

President and CEO

October 29, 2021

Forward-Looking Statements



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5-Year Business Plan (FY2021-FY2025) for Sustainable Growth



Under ESG management, we will realize our 2025 Goal, Global Pharma Innovator with Competitive Advantage in Oncology, and will shift to further growth toward our 2030 Vision



2030 Vision

Innovative Global
Healthcare Company
Contributing to the
Sustainable Development
of Society

- ♦ Global top 10 in Oncology
- Additional growth pillars being source of revenue and profit
- ♦ New products being source of profit in each business unit
- Contributing to sustainable development of society through our business

As of FY2020

- Oncology business launched
- Edoxaban growing
- Regional value being enhanced
- ◆ AZ strategic alliance
- Increased RD investment

Strategic Pillars for the 5-Year Business Plan (FY2021-FY2025)



Realize 2025 Goal and Shift to Further Growth

FY2025 Financial Targets

- ◆ Revenue: 1.6 Tn JPY (Oncology > 600.0 Bn JPY)
- ◆ Core Operating Profit Ratio before R&D Expense: 40%

- **♦** ROE > 16%
- **♦ DOE* > 8%**

Maximize 3ADCs

- Maximize ENHERTU and Dato-DXd through strategic alliance with AstraZeneca
- Maximize HER3-DXd without a partner
- Expand work force and supply capacity flexibly depending on changes around product potential

Profit growth for current business and products

- Maximize Lixiana profit
- Grow Tarlige, Nilemdo, etc. quickly
- Transform to profit structure focused on patented drugs
- Profit growth for American Regent and Daiichi Sankyo Healthcare

Identify and build pillars for further growth

- Identify new growth drivers following 3ADCs
- Select and advance promising post DXd-ADC modalities

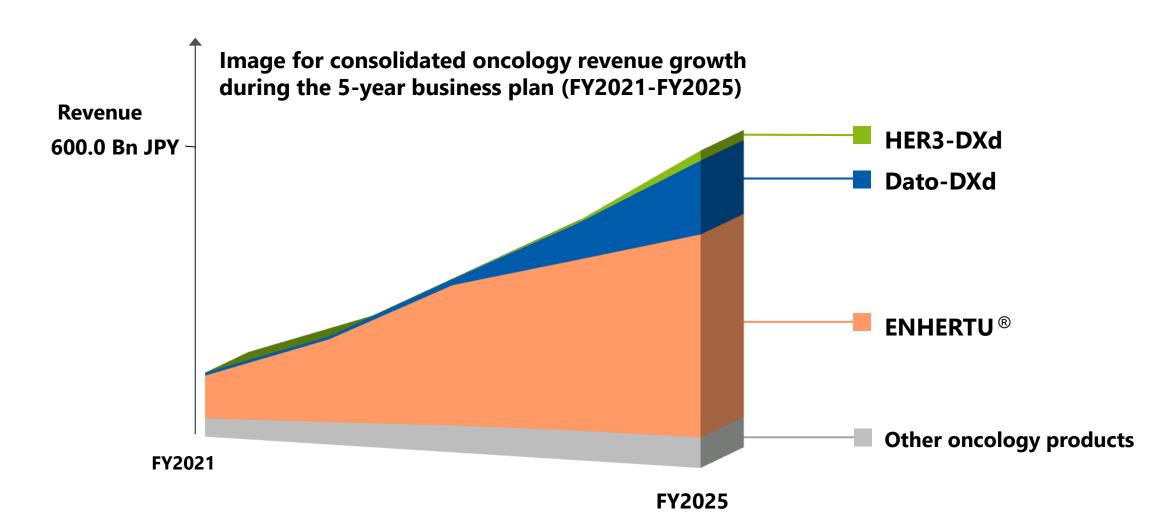
Create shared value with stakeholders

- Patients: Contributing to patients through "Patient Centric Mindset"
- Shareholders: Balanced investment for growth and shareholder returns
- Society: Environment load reduction across the value chain, and actions against pandemic risks
- Employees: Create one DS culture through fostering our core behaviors
- Data-driven management through DX, and company-wide transformation through advanced digital technology
- **♦** Agile decision making through new global management structure

Oncology Revenue Target



Targeting > 600.0 Bn JPY in FY2025 by maximizing 3ADCs



ENHERTU®: Clinical Development Plan | Breast cancer



As of Sep 2021			FY2020	FY2021	FY2022	Planning
		Metastatic 3L~	DESTINY-Breast01 complete	d		
			DESTINY-			
		Metastatic 2L	DESTINY-Breast03 monoth	erapy vs T-DM1		
HER2 Po	sitive	Metastatic 1L				
		Post-neoadjuvant				
		Neoadjuvant				Phase 3
		Adjuvant				Phase 3
	HR+ HR-	Metastatic Post Chemo	DESTINY-Breast0			
		Post-neoadjuvant				Phase 3
HER2 Low	HR+	Metastatic Chemo Naive	D			
		Metastatic Endocrine Therapy				Phase 3
	HR-	Metastatic 1L	BEGONI			
		Neoadjuvant				Phase 3

Study initiation & end points are all shown as either beginning of 1H or 2H PC: physician's choice

Ph 3 ongoing

New

Ph 1 ongoing

ENHERTU®: Clinical Development Plan | Gastric cancer & NSCLC



As of Sep	o 2021		FY2020	FY2021	FY2022	Planning
Gastric	HER2 Positive	Advanced/ Metastatic 3L~	DESTINY-Gastric01			
		Advanced/ Metastatic 2L	DESTINY-Gastric02 monot			
				DESTINY-Gastrio	c04 mono vs ramucirumab+paclitaxe	
		A diverse and /	DESTIN			
		Advanced/ Metastatic 1L				Phase 3
	HER2 Expressing	Advanced/ Metastatic 2L~	DESTINY-Lung01 mor	notherapy		
		Advanced/ Metastatic 2L				Phase 3
		Advanced/ Metastatic 1L				
NSCLC						Phase 3
	HER2 Mutated	Advanced/ Metastatic 2L~	DESTINY-Lung01 mor	notherapy		
		Advanced/ Metastatic 1L				
	Expressing /Mutated	Early disease				Phase 3

Study initiation & end points are all shown as either beginning of 1H or 2H

Ph 3 ongoing

New

Ph 1 ongoing

NSCLC: non small cell lung cancer

ENHERTU®: Clinical Development Plan | CRC & other tumors



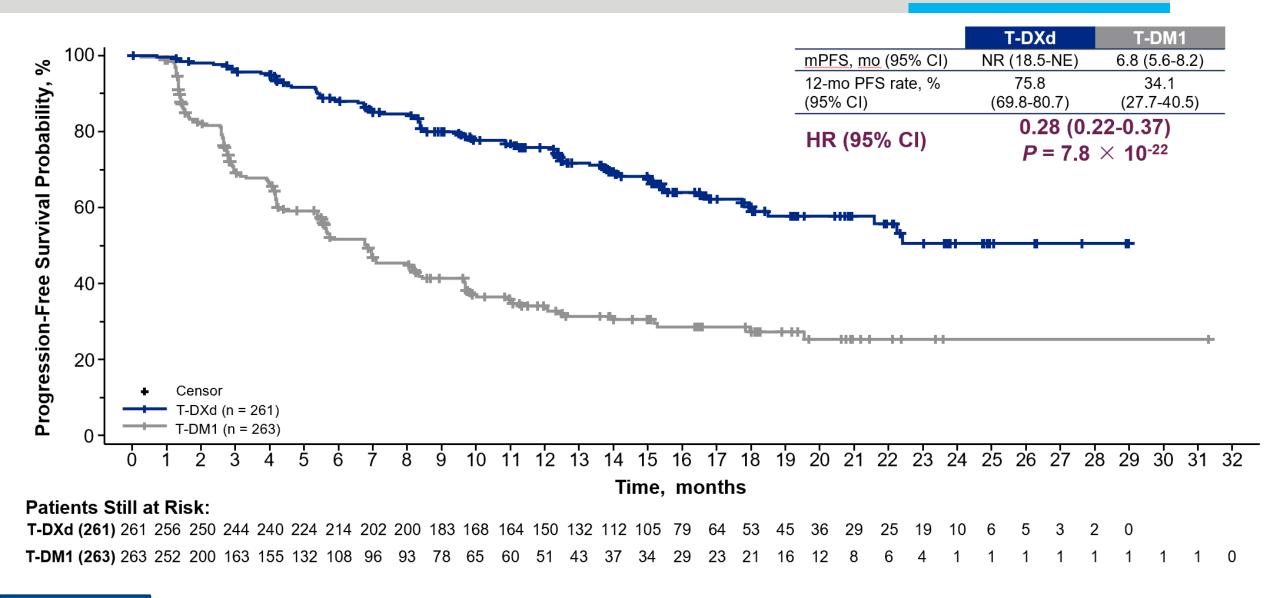
As of Sept	ember 2021		FY2020		FY2021		FY2022		Planning
CRC	HER2 Express ing	Metastatic 3L	DESTINY-CRC01	1 monotherapy DESTINY-CRC02 monotherapy					
		Metastatic 2L							Phase 3
		Metastatic 1L							Phase 3
	HER2 Express ing	∣ າ ເ	Nivolumab combination (breast, bladder)						
Other			Pembrolizumab combination (breast, NSCLC)						
Tumors/ multiple					DES	TINY-PanTumo	r02		
tumors		Ovarian							Phase 2
	HER2 Mutated	Metastatic 2L	DESTINY-PanTumor01				r01		
Ph 1 ongoing	Ph 2 ongo	ing Ph 3 ongo	ping New	Completed)				

Study initiation & end points are all shown as either beginning of 1H or 2H

CRC: colorectal cancer, NSCLC: non small cell lung cancer

Primary Endpoint: PFS by BICR

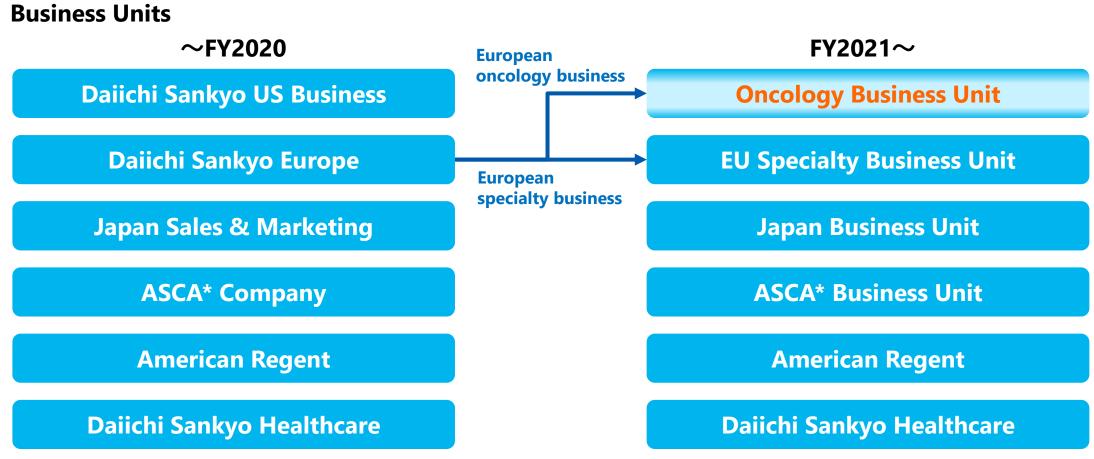




Creation of Oncology Business Unit



Oncology Business Unit was created to align US and European oncology businesses as well as global oncology business functions under one team to respond to the rapid changes we see in standards of care, and the oncology market



*Asia, South and Central America

Passion for Innovation.
Compassion for Patients.™

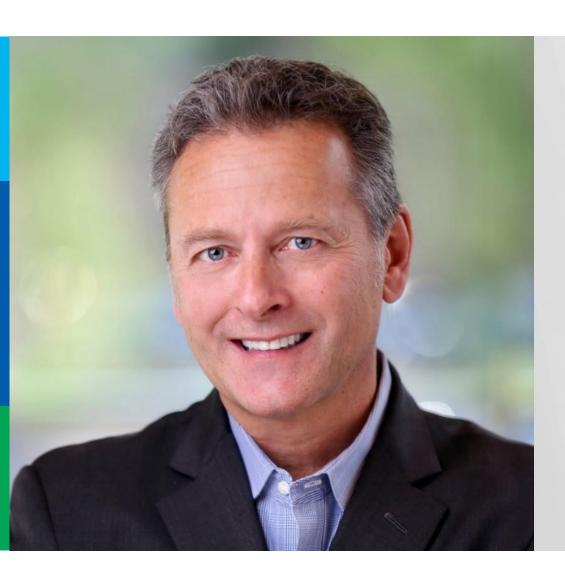


ENHERTU® Business Briefing

Ken Keller Head of Oncology Business Unit October 29, 2021

Daiichi Sankyo OBU Global Leadership





Ken Keller

Global Head, Oncology Business President and CEO, Daiichi Sankyo, Inc.

- Joined Daiichi Sankyo in 2014
- Revamped U.S. business structure to focus on multiple oncology launches including ENHERTU[®] as part of Daiichi Sankyo's 2025 Goal
- More than 30 years of experience in the pharmaceutical industry including 22 years at Amgen
- Held senior regional and global leaderships roles supporting major biologics including Aranesp, Enbrel, Neulasta, Neupogen, Prolia, Vectibex and Xgeva

Daiichi Sankyo OBU Global Leadership Team





US EAST COAST

JAPAN



Mary Pinder-Schenck

Vice President, Head of Global Oncology Medical Affairs



Ken Keller

President and CEO, Daiichi Sankyo, Inc.

Global Head of Oncology Business



Dan Switzer

Head of US Oncology Business Division



Nagatomo Hamahata

Global Head of Oncology Alliance Management



Rich Jones

Executive Director, Global Oncology Business Strategy & Analysis



Nadine Sprangers

Vice President, Head of Global Oncology Market Access & Pricing



Markus Kosch

Head of EU Oncology Business Division



Kenji Shigeta

Head of Global Oncology Marketing

Daiichi Sankyo's Transformation into Global Oncology Leader



POTENTIAL FOR ~12 APPROVALS FROM DXd ADC PORTFOLIO IN MULTIPLE INDICATIONS ACROSS MORE THAN 30 COUNTRIES IN NEXT 5 YEARS

HER2 ADC TROP2 ADC HER3 ADC B7-H3 ADC HERTHENA TROPION **DESTINY Trastuzumab Deruxtecan Datopotamab Deruxtecan Patritumab Deruxtecan** (Dato-DXd) (HER3-DXd) (T-DXd) **DS-7300** mNSCLC with and without EGFR mutant mNSCLC • SCLC HER2+ mBC & eBC **AGAs** HER2 low mBC mBC Esophageal squamous TNBC cell carcinoma mCRC HER2+ mGC HR+ / HER2- mBC Castration resistant HER2 mutant mNSCLC prostate cancer 4 U.S. FDA BTDs have been granted so far AstraZeneca Daiichi-Sankyo COMBINATION STRATGIES WITH IMMUNOTHERAPY, OTHER ANTI-CANCER MEDICINES

COMBINATION STRATGIES WITH IMMUNOTHERAPY, OTHER ANTI-CANCER MEDICINES
PART OF CLINCIAL DEVELOPMENT PROGRAMS ACROSS PORTFOLIO

ENHERTU®: Roadmap to Transforming HER2 Targetable Cancers



Establish Foundation

Achieve 3L HER2+ mBC and 2L/3L HER2+ mGC market leadership in all launch countries



Build Market Leadership

Establish ENHERTU as HER2 medicine of choice for 2L HER2+ mBC and become market leader worldwide

#1

Redefine HER2 Treatment Paradigm

Rethink HER2 targetable population with potential of ENHERTU efficacy in HER2 low mBC and HER2 mutant mNSCLC



Elevate Outcome Expectations

Push new boundaries with further development in neoadjuvant/adjuvant eBC, 1L mBC, combination strategies and other cancers

2020 - 2021

2022

2022 - 2023

2024 +

ENHERTU®: Solid Performance Globally



ENHERTU #1 IN 3L HER2+ MBC IN EVERY COUNTRY FULLY LAUNCHED APPROVED IN 37 COUNTRIES | ~8,000 PATIENTS TREATED WORLDWIDE

US

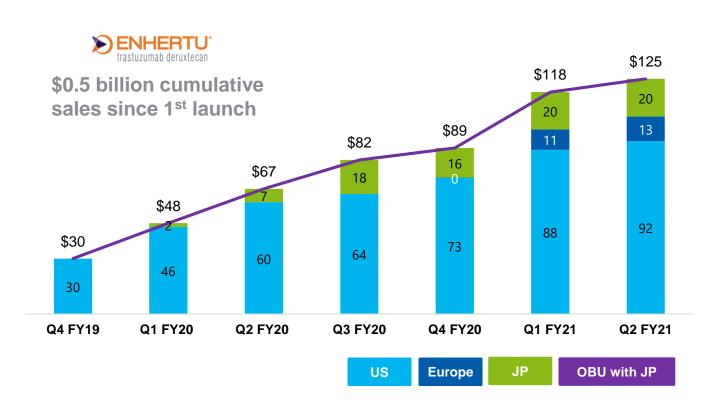
- #1 3L HER2+ mBC medicine
- Achieved >40% market share in 3L
- Prescriber research shows ~65% of patients will receive ENHERTU in 3L+
- Rapid adoption in HER2+Gastric
- ~ 5,000 patients treated
- Preparations in place for 2L approval

Europe

- #1 3L HER2+ mBC medicine in France & UK
- Launched in Austria, Denmark, Finland, France, Luxembourg, Norway, Sweden & UK – with more countries to join soon
- Demand sales +31% in August 2021 vs July
- ~ 1,000 patients treated

JP

- #1 3L HER2+ mBC & 3L HER2+ mGC medicine
- 30% and 45% market share respectively
- ~ 2,100 patients treated in total
- Well managed ILD risk with BC & GC HCPs



3L HER2+ mBC approvals: Australia, Brazil, Canada, Israel, Japan, UK and U.S.

2L/3L HER2+ mGC approvals: Israel, US

3L HER2+ mGC approvals: Japan

ESMO 2021: Defining Moment for Daiichi Sankyo's Transformation into Global Oncology Leader



18 Abstracts with 6 Oral Presentations

- 1 Presidential Symposia
- 4 Late-Breaking Presentations
- 3 Preferred Papers
- 2 Mini Oral Presentations
- 9 Posters
- 1 NEJM Publication

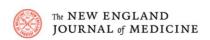
4 Late-Breaking Presentations











ORIGINAL ARTICLE

Trastuzumab Deruxtecan in HER2-Mutant Non-Small-Cell Lung Cancer

Bob T, Li, M.D., Ph.D., M.P.H., Egbert F. Smit, M.D., Ph.D.,
Yasushi Goto, M.D., Ph.D., Kazuhiko Nakagawa, M.D., Hibiki Udagawa, M.D.,
Julien Maziřers, M.D., Misako Nagasaka, M.D., Ph.D.,
Lyudmila Bazhenova, M.D., Andreas N. Saltos, M.D.,
Enriqueta Felip, M.D., Ph.D., Jose M. Pacheco, M.D., Maurice Pérol, M.D.,
Luis Paz-Ares, M.D., Kapil Saxena, M.D., Ryota Shiga, B.Sc.,
Yingkai Cheng, M.D., Ph.D., Suddhasatta Acharyya, Ph.D.,
Patrik Vitazka, M.D., Ph.D., Javad Shahidi, M.D., David Planchard, M.D., Ph.D.,
and Pasi A, Jänne, M.D., Ph.D., for the DESTINY-Lungol Trial Investigators*

Additional Data



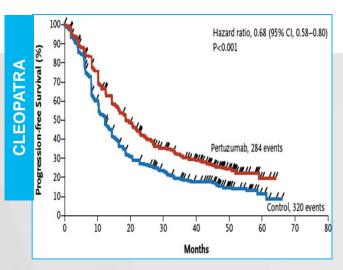


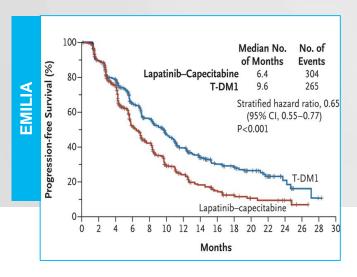
DS-7300
4th DXd ADC in Clinical Development

ENHERTU®: New Standard of Care for 2L mBC

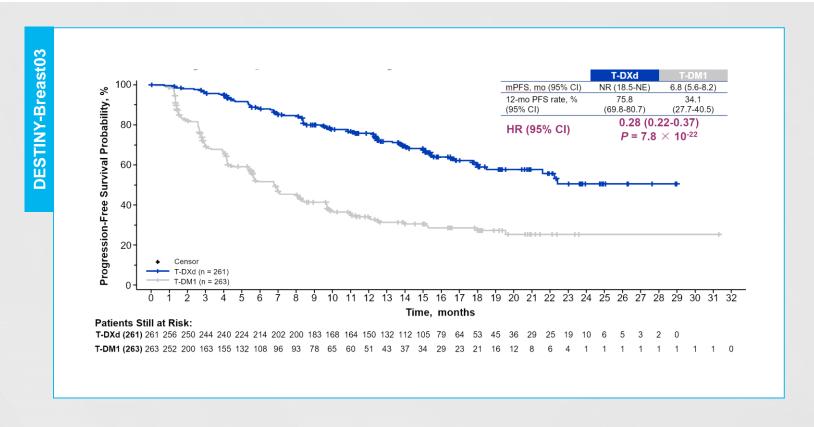


PRIOR ADVANCES IN HER2+ MBC





DESTINY-Breast03 Primary Endpoint: PFS by BICR

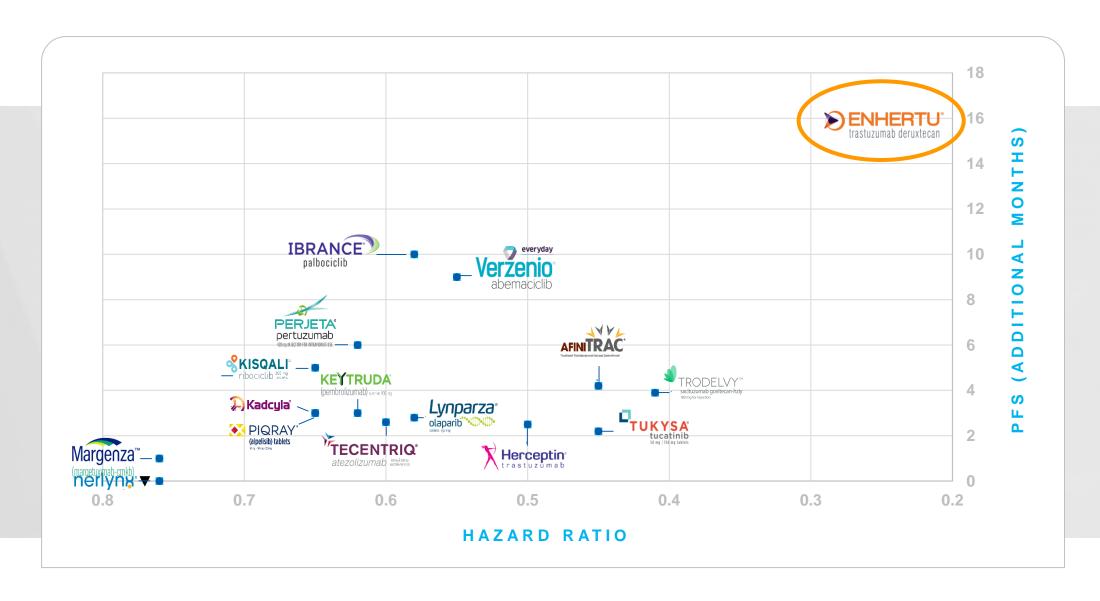


Cortes J, et al. ESMO 2021

Median PFS follow-up for T-DXd was 15.5 months (range, 15.1-16.6) and for T-DM1 was 13.9 months (range, 11.8-15.1) HR, hazard ratio; INV, investigator; mo, month; NE, not estimable; NR, not reached.

ENHERTU®: Historical Unprecedented Efficacy In Breast Cancer





ENHERTU®: Media Reaction to DESTINY-Breast03





With Massive Tumor Progression Edge, AstraZeneca-Daiichi's Enhertu Shows Roche's Kadcyla Who's the Better Breast Cancer Drug

"The showing is so impressive that the study authors concluded that the study, dubbed Destiny-Breast03, will lead to a paradigm shift in the treatment of HER2-positive breast cancer."

ENDPOINTS NEWS

AstraZeneca, Daiichi Sankyo's ADC Enhertu Blows Away Roche's Kadcyla in Second-line Advanced Breast Cancer

"Getting into earlier patients is now the goal, starting with Enhertu's complete walkover of a Roche drug in second-line breast..."

MEDPAGE TODAY

Trastuzumab Deruxtecan: New Standard of Care in Pretreated, Advanced Breast Cancer?

"These PFS curves from DESTINY03 are absolutely startling...I don't believe I've seen a hazard ratio like this in HER2 breast cancer before."

- Dr. Shanu Modi, Memorial Sloan Kettering



Biotech Strategy Blog

"...the last time I saw similar shock and awe or such fevered excitement in breast cancer was... a decade ago...so it has been a long time coming for a new standard of care to be seen in this disease. This is what we live for in oncology, records and standards are meant to be broken."

Sally Church, Blog Editor



Drug Combination Shows Revolutionary Results in Extending Life for Tough-to-treat Breast Cancer

- "...this potent antibody drug...will dramatically change the treatment for HER2 positive breast cancer."
- Dr. Sara Tolaney, Dana-Farber Cancer Institute

ENHERTU®: KEE Reaction to DESTINY-Breast03



COMMON THEME: ACCELERATE RESEARCH FOR USE OF ENHERTU IN EARLIER LINES OF HER2+ BREAST CANCER

"Super impressive outcome.

T-DXd is the winner. I agree that the next question is, what do we do with the T-DM1."

"I think that for a majority of patients and for physicians like myself who treat metastatic HER2 positive breast cancer, it's going to lead to a paradigm shift in how we treat this patient population." "It was a really substantial difference in the two treatment arms. **This** data is nothing short of phenomenal and will be practice changing." "The world is brighter for women with HER2 overexpressing breast cancers."

"I don't believe I've seen a hazard ratio like this in HER2 breast cancer before." "A drug like this has potential to do better, much better than a 5-6% cure rate in metastatic breast cancer. The MOA and uniqueness of the payload could move the cure rate up for metastatic disease considerably. Given the reassuring safety, this drug should move to curative early breast cancer setting as soon as possible. The only competitor is tucatinib plus trastuzumab in early breast cancer as a well-tolerated regimen – "two gunslingers to meet on the streets."

"Ushering in a new standard of care for the second line treatment of metastatic HER2 positive breast cancer—
trastuzumab deruxtecan dramatically outperforms T-DM1
with acceptable toxicity"

"ENHERTU was the star of the show"

"We've all seen the little steps we've been taking, and **this is a much bigger step**... kudos to DS and AZ and the vision of everyone going forward with this drug...."

"How quickly could we test this in early breast cancer? We are highly interested to investigate this drug in early breast cancer..." "Given this impressive magnitude of benefit, I believe we will be able to eliminate breast cancer as cause of death...'

"Jaw dropping waterfall plot"

"This is the confirmation of our clinical impression – that is drug will **change the way we treat breast cancer**, not just HER2 positive breast cancer...."

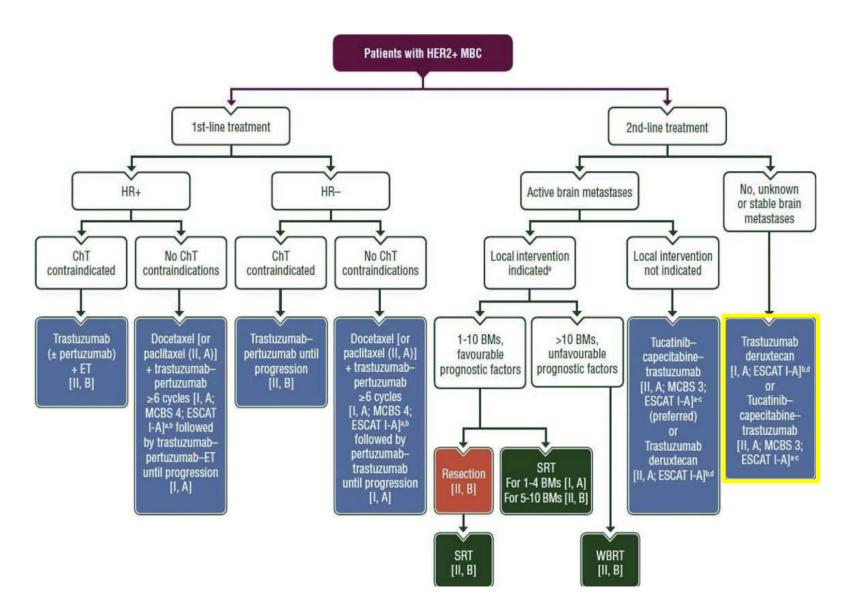
"ADCs will be the pillar of treatment in 5 years for all chemo-sensitive disease....we are going to substitute chemotherapy for ADCs now..." "This potent antibody drug...will dramatically change the treatment for HER-2 positive breast cancer."

"It was a really substantial difference in the two treatment arms.

T-DM1 is a good adjuvant drug, but we are always striving to do better. ENHERTU is offering that better option."

Annals of Oncology Published Oct 19, 2021: ESMO Clinical Practice Guideline for the diagnosis, staging and treatment of patients with metastatic breast cancer, supports **ENHERTU®** as the New standard of care



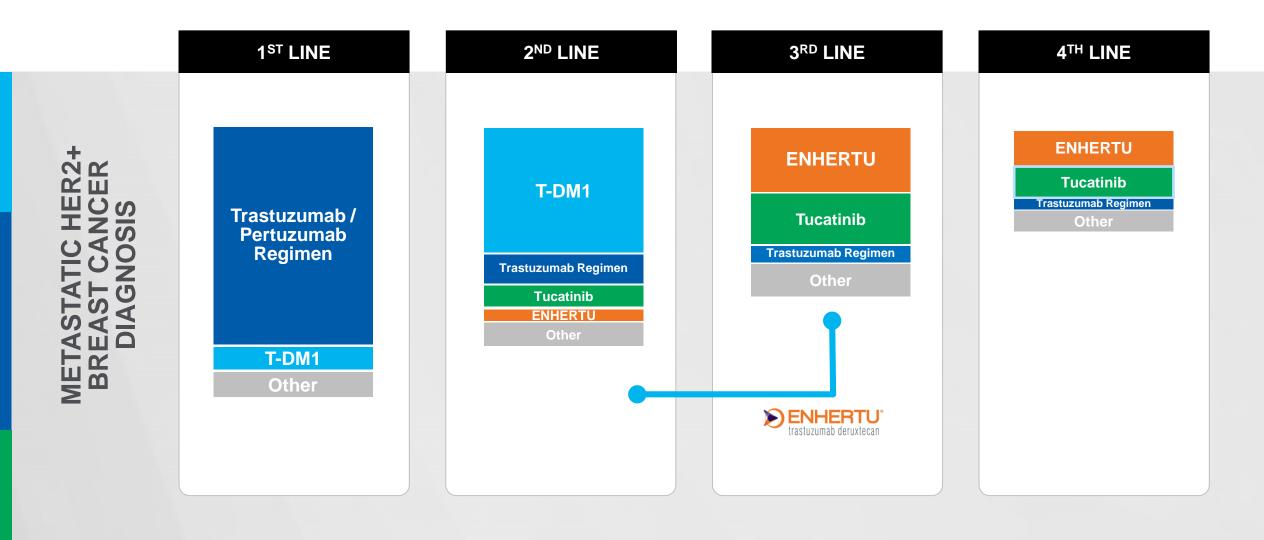


Second line language in ESMO Guideline

- Based on the strength of these efficacy and safety data, it is reasonable to consider trastuzumab deruxtecan the new standard second-line therapy in regions where this drug is available [I, A], moving T-DM1 to a later-line setting.
- Trastuzumab deruxtecan should be given as second-line therapy after progression on a taxane and trastuzumab [I, A].
- T-DM1 is a second-line treatment option after progression on a taxane and trastuzumab in cases where trastuzumab deruxtecan is not available [I, A; ESMO-MCBS v1.1 score: 4; ESCAT score: I-A].
- Tucatinib/capecitabine/trastuzumab or trastuzumab deruxtecan may be used in the second-line setting in selected patients with BMs [II, A]

ENHERTU®: Opportunity to Displace T-DM1 in mBC





Sources: ENHERTU HER2+ ATU and PCA (ZoomRx); internal sources

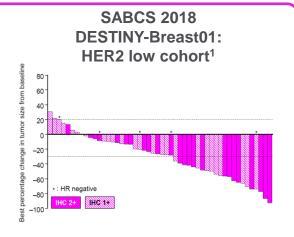
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ENHERTU® Next: Transform Treatment Landscape for Previously "Un-targetable" HER2 Low

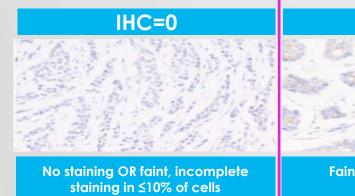


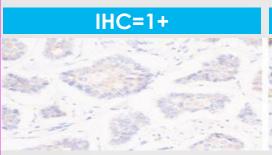
CURRENT SITUATION

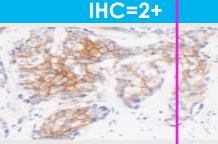
- Many HER2 targeted therapies have been approved in HER2+ segment
- No HER2 targeted therapy has demonstrated meaningful efficacy beyond HER 2+ population
- In HR+ segment, ET and CDK4/6 treatment is standard of care and progressing patients receive chemotherapy with limited success
- DESTINY-Breast04 and DESTINY-Breast06 trials targeting HER2 Low patients is projected to read out in Q4 FY2021 for DESTINY-Breast04, and additional indication from DESTINY-Breast06 is expected to contribute commercially to the current 5-year business plan
- HER2 testing is well entrenched across markets (>95% in mBC)
- HER2 Low patients can be identified with the current IHC SOC assays

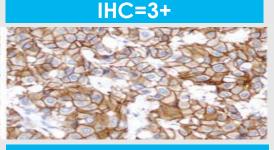


1 Modi et al, SABCS, 2018; Poster # P6-17-02, Abstract #486









Faint, incomplete staining in >10% of cells

Weak – moderate, complete staining in >10% of cells

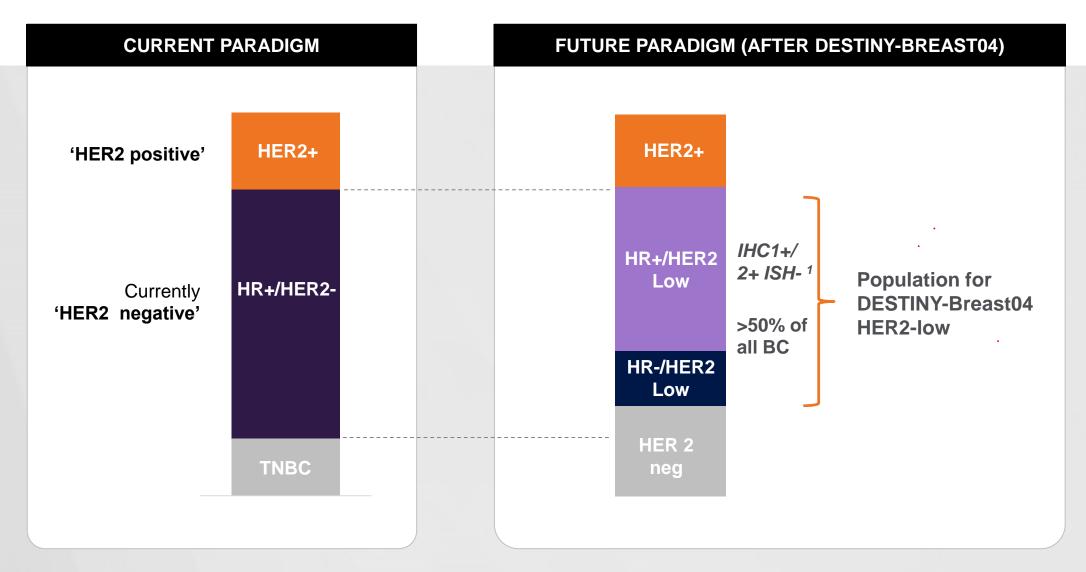
Intense - complete staining in >10% of cells

"HER2 Low" - IHC1+/2+, ISH-HER2 targeted drugs have tried and all failed "HER2 +" - IHC2+/3+, ISH+ trastuzumab, pertuzumab, T-DM1, neratinib, tucatinib, lapatinib

ENHERTU®: DESTINY-Breast04 and DESTINY-Breast06



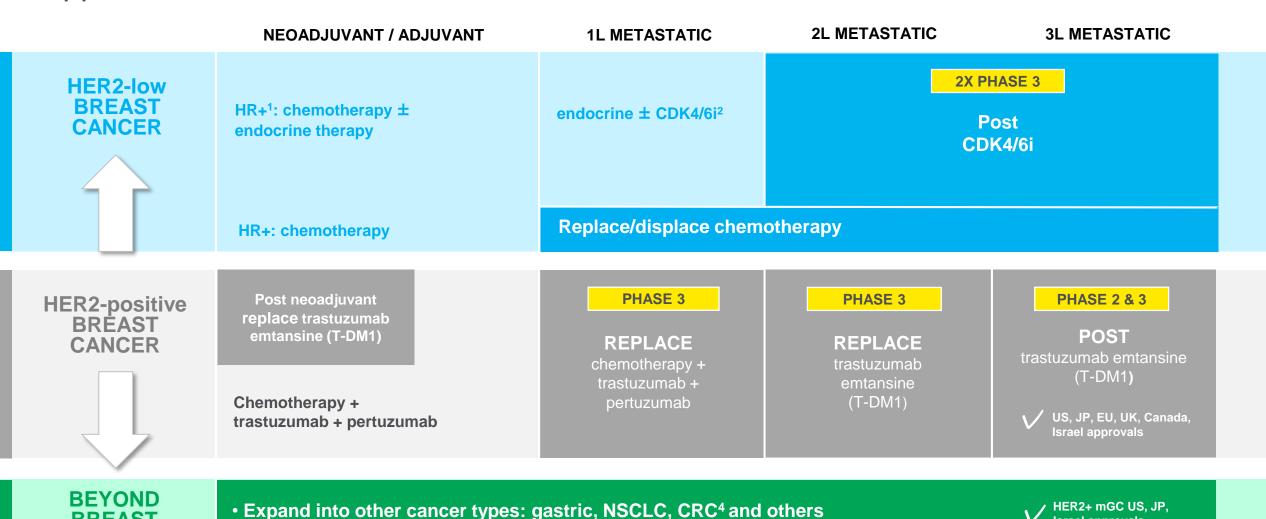
to Potentially Change Treatment Paradigm for Large Segment of Breast Cancer



ENHERTU®: Clinical Development Program Highlights



Opportunities across breast cancer, HER2-low and other tumors



• Conducting multiple combination trials to push the boundaries of patient outcomes

BREAST

CANCER

Israel approvals

Dato-DXd: Potentially Best-in-Class TROP2 Directed ADC in NSCLC and Breast Cancer



TROP2

is a widely expressed in solid tumors, including high expression in NSCLC where it is associated with poor prognosis

Initiated
Global
Phase 3
TROPIONLung01⁴ vs
docetaxel in
2L, 3L mNSCLC
without AGAs
post IO and PBC

Additional Phase 1/2 Trials in NSCLC

- TROPION-Lung05⁵ monotherapy in 3L+ mNSCLC with AGAs
- TROPION-Lung02⁶ & TROPION-Lung04⁷
 IO combos
 (pembrolizumab, durvalumab with or without PBC) in 1L, 2L, 3L mNSCLC without AGAs

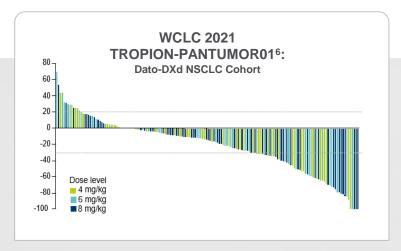
Phase 1 in TNBC & HR+/HER2- mBC

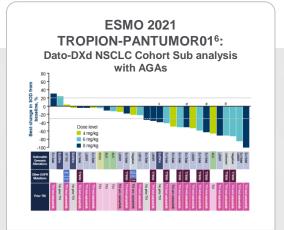
Promising results in TNBC cohort from TROPION-PanTumor018

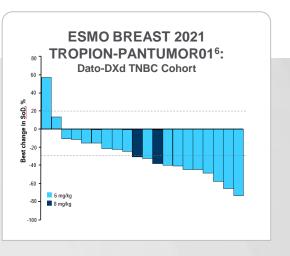
AGAs: Actionable Genomic Alterations; IO: Immunotherapy; ORR: Objective Response Rate; OS: Overall Survival; PBC: Platinumbased Chemotherapy;

Salvage Chemo in NSCLC

limited efficacy (ORR of 5-23% and OS of 5.7-12.6 months) post IO and PBC^{1,2,3}







^{1.} Hotta K, et al. *J Thorac Oncol.* 2007. 2. Rothschild SI, et al. *ESMO Open.* 2021. 3. Garon E, et al. *The Lancet* 2014 4. NCT04656652 5. NCT04484142 6. NCT04526691 7. NCT04612751 8. NCT03401385

HER3-DXd: First-in-Class HER3 Directed ADC in NSCLC and Breast Cancer

1/2 Trial³

in NSCLC

osimertinib

combo in 1L, 2L

mNSCLC with



HER3

Is frequently overexpressed in NSCLC as well as other types of solid tumors

Salvage

limited efficacy (PFS of 2.8-3.2 months) post **EGFR TKIs** and PBC¹

Initiated Global Phase 2 **HERTHENA-**Lung01²

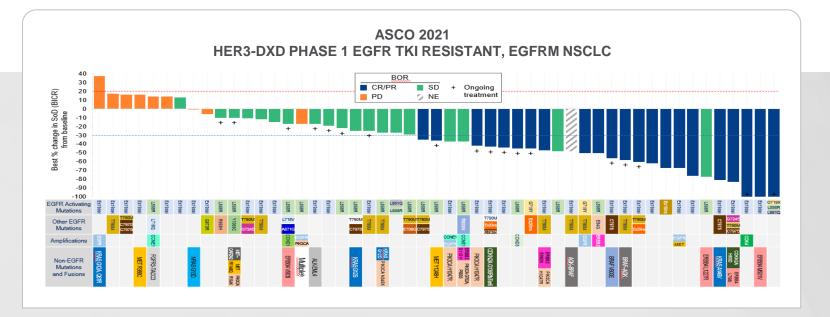
in 3L+ EGFR mutant mNSCLC post osimertinib and PBC

Additional Breast Cancer & Beyond Phase

• Phase 1/2⁴ in HR+/HER2mBC and TNBC across lines of therapy Phase 2⁵ in colorectal cancer

EGFR: Epidermal Growth Factor Receptor; IO: Immunotherapy; PBC: Platinum-based Chemotherapy: PFS: Progression-Free Survival TKI: Tyrosine

Chemo



^{1.} Yang, CJ. BMC Pharmacol Toxicol. 2017.

^{2.} NCT04619004

^{3.} NCT04676477

^{4.} NCT02980341

^{5.} NCT04479436

ENHERTU®, Dato-DXd & HER3-DXd: Lung Cancer Clinical Development Highlights









1L METASTATIC 2L METASTATIC 3L METASTATIC

NSCLC with AGAs* ~49%

- EGFRm ~17%
- HER2m ~2-4%

*AGA= actionable genomic mutations defined as % of non-squamous NSCLC patients who are positive for EGFRm (excluding exon 20 deletion), HER2m, ALK, ROS1, NTRK1, BRAF, KRAS G12C, Met, Ret HER3-DXd
Phase 1 (1L, 2L EGFRm NSCLC)
HER3-DXd combo with osimertinib

HERTHENA-Lung01
Phase 2 (3L EGFRm NSCLC)
HER3-DXd monotherapy

TROPION-Lung05
Phase 2 (3L+ NSCLC with AGA)
Dato-DXd monotherapy

DESTINY-Lung01
Phase 2 (2L HER2m and HER2+ NSCLC)
ENHERTU Monotherapy

DESTINY-Lung02
Phase 2 (2L HER2m NSCLC)
ENHERTU Monotherapy Post PBC

NSCLC without AGAs** ~51%

** TL01 and TL08 does not exclude patients who have KRASG12C mutations in markets where KRASG12C inhibitors are not approved TROPION-Lung02
Phase 1b (1L, 2L, 3L NSCLC without AGAs)
Dato-DXd combo with pembrolizumab with or without PBC

TROPION-Lung04
Phase 1b (1L, 2L, 3L NSCLC without AGAs)
Dato-DXd combo with durvalumab with or without PBC

TROPION-Lung08
Phase 3 (1L NSCLC without AGA)
Dato-DXd + pembrolizumab vs. pembrolizumab

TROPION-Lung01
Phase 3 (2L, 3L NSCLC without AGA)
Dato-DXd vs. docetaxel

DESTINY-Lung03
Phase 1b (1L, 2L, 3L HER2+ NSCLC)
ENHERTU combo with durvalumab and chemotherapy

By 2030 DS will be a Global Innovator, ADC Leader Across Multiple Tumor Types with a Competitive Advantage in Oncology



FY2021 - FY2025

As of FY2021

- · Oncology Business Unit launched
- ENHERTU[®] approved globally in mBC and mGC
 - ENHERTU[®] granted 4th
 Breakthrough Therapy
 Designation in the US based
 on DESTNY-Breast03 results
 - Well-positioned to be market leader in 2L HER2+ mBC
 - Positioned to transform HER2 Low treatment paradigm
- Multiple drugs in breast and lung cancer launched

5-YEAR BUSINESS PLAN (FY2021-FY2025)

REALIZE
2025 GOAL
AND SHIFT
TO FURTHER
GROWTH

2030 VISION

Innovative Oncology Business Contributing to the Sustainable Development of Society

- Global Top 10 Oncology Company
- Leader in ADC technology across multiple tumors types
- ENHERTU®, Dato-DXd and HER3-DXd achieve blockbuster status
- Complemented by promising ADC candidate DS-7300
- DS recognized as trusted and respected Oncology business across the globe

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